

July 20, 2001

Docket No. 98N-0337 Food and Drug Administration 5630 Fishers Lane, Room 1061 Rockville, MD 20852 7650 °01 JLL 20 A10:28

APPLICATION FOR EXEMPTION

Subject:

Minoxidil Topical Solution USP, 5% (For Men)

ANDA 75-598

Docket No. 98N-0337 APPLICATION FOR EXEMPTION

## **Statement of Purpose**

Pursuant to 21 CFR 201.66(e), Perrigo requests an exemption from 21 CFR 201.66(c) and (d) in the form of a temporary deferral of the implementation of the requirements of this regulation. This deferral is requested because there is not currently approved labeling in the Drug Facts format for the reference listed drug available to the Perrigo Company. The exemption would apply to all current and future SKUs of the drug product.

The reference listed drug for this ANDA is Rogaine® Extra Strength (For Men) (NDA 20-834).

## **Background of the Request**

From the time that the final rule was issued in 1999, it has been the understanding of the Perrigo Company, through several contacts with the Office of Generic Drugs, that the Agency would not approve ANDA labeling formatted according to the requirements described in 21 CFR 201.66 until approved reference listed drug labeling similarly formatted was available. Perrigo further understands, based on these contacts, that in the absence of approved reference listed drug labeling in drug facts format, ANDA labeling could not be converted regardless of the May 2002 deadline.

We believe that it is the Office of Generic Drugs' position that Drug Facts and non-Drug Facts format labeling may not be 'the same' as required by the Food Drug and Cosmetic Act under part 505 (j)(2)(A), and in fact, that the ANDA holder cannot know if the labeling will be 'the same' until the reference listed drug labeling is available for comparison. Therefore, in order to ensure continuing compliance with both the statue and the regulation, a temporary deferral of the implementation date is required until approved reference listed drug labeling is available in Drug Facts format.

96N-0337

515 Eastern Avenue Allegan, Michigan 49010 (616) 673-8451 APP27

In a letter from Dr. Charles Ganley to the Consumer Healthcare Products Association dated August 9, 1999, it was recommended that ANDA holders submit a request for deferral in those cases where the reference listed drug has not received approval for labeling in the Drug Facts format in sufficient time to allow conversion of the ANDA product labeling by the regulatory compliance date.

### "Templates" for Drug Facts Labeling

The Office of Generic Drugs has published in a February 2001 draft guidance, certain templates for drug facts labeling of particular drugs, and has since published additional templates for products for which there is not approved reference listed drug (RLD) labeling in Drug Facts format. The February 2001 draft guidance also made reference to the potential for ANDA applicants to submit changes to implement Drug Facts labeling in the absence of an approved reference listed drug in this format.

Our discussions as late as July 2001 with OGD representatives have verified that the presence of a published template does not confer any special status to a drug product in the absence of approved RLD labeling. OGD will not grant approval for a supplement to implement drug facts labeling for an OTC ANDA product before the approval of the RLD in the same format. Further, since labeling in drug facts format and non-drug facts format is not considered to be "the same", ANDA holders may not implement Drug Facts format labeling by way of an annual report. The potential finalization date and content of the February 2001 draft guidance is unknown.

#### Drugs Approved After April 16, 1999

The final rule, published on March 17, 1999, states that OTC ANDA drug products approved after April 16, 1999, must meet the requirements of part 201.66 immediately upon approval. However, the Office of Generic drugs has continued to approve ANDAs after April 16, 1999, without labeling that complies with section 201.66 and without comment as to the need to meet the requirements.

This drug product was approved on June 13, 2001. However, as of the date of this request, there is not approved RLD labeling in the Drug Facts format available to Perrigo.

According to our discussions with the Office of Generic Drugs, in the case where an ANDA for an OTC drug product is approved after April 16, 1999, without Drug Facts labeling, that product may be marketed with labeling approved in the ANDA. When approved reference listed drug labeling in Drug Facts format becomes available to the ANDA applicant, the product labeling should then be updated with appropriate notice to the application.

## Length of the Deferral Request

Due to the large number of store-brand private labels maintained by Perrigo for each ANDA OTC drug product, converting the labeling to Drug Facts format requires significant time and resources. For any drug product for which Drug Facts format labeling is not available as of the date of this letter, Perrigo is submitting a request for a temporary deferral of implementation.

At the time that approved Drug Facts format labeling becomes available for each RLD, Perrigo will immediately act to file a Changes Being Effected Supplement for approval of the new labeling in the relevant ANDA. The product will then be entered into our labeling conversion schedule. Due the length of time required to prepare labeling, submit a CBE supplement, and finally convert the labeling of a product, we anticipate that conversion for a particular product can be accomplished within approximately six months from the approval of the labeling supplement or twelve months from when the RLD labeling is first approved and available to Perrigo in Drug Facts format.

If the reference listed drug for this ANDA has approved labeling available in Drug Facts format by the compliance date of May 2002, then this deferral is not anticipated to be required beyond May 2003.

If there are any questions concerning this request, please contact me by phone at (616) 673-9745 or fax at (616) 673-7655. Thank you for your attention to this matter.

Sincerely,

L. PERRIGO COMPANY

Brian Schuster

Manager, ANDA Submissions

CC:

Gary Buehler, Director
Office of Generic Drugs
FDA/CDER
Metro Park North II
7500 Standish Place, Room 150
Rockville, MD 20855

#### DEPARTMENT OF HEALTH AND HUMAN SERVICES

FOOD AND DRUG ADMINISTRATION

# APPLICATION TO MARKET A NEW DRUG, BIOLOGIC, OR AN ANTIBIOTIC DRUG FOR HUMAN USE

(Title 21, Code of Federal Regulations, 314 & 601)

Form Approved: OMB No. 0910-0338 Expiration Date: March 31, 2003 See OMB Statement on page 2.

FOR FDA USE ONLY

APPLICATION NUMBER

APPLICANT INFORMATION									
NAME OF APPLICANT L. Perrigo Company		JUL 19 2001							
TELEPHONE NO. (Include Area Code) 616- 673-8451	FACSIMILE (FAX) Number (Include Area Code) 616- 673-7655								
APPLICANT ADDRESS (Number, Street, City, Ste and U.S. License number if previously Issued): 515 Eastern Ave. Allegan, MI 49010	te, Country, ZIP Code or Mail Code,	AUTHORIZED U.S. AGENT NAME & ADDRESS (Number, Street, City, State, ZIP Code, telephone & FAX number) IF APPLICABLE							
PRODUCT DESCRIPTION		<del></del>							
NEW DRUG OR ANTIBIOTIC APPLICATION NUMBER, OR BIOLOGICS LICENSE APPLICATION NUMBER (If previously issued) 75-598									
ESTABLISHED NAME (e.g., Proper name, US Minoxidil Solution	STABLISHED NAME (e.g., Proper name, USP/USAN name)  PROPRIETARY NAME (trade name) IF ANY								
CHEMICAL/BIOCHEMICAL/BLOOD PRODUC 2,4-Pyrimidinediamine, 6-(1-piperidinyl)-3	T NAME ( <i>If any</i> ) -oxide.	CODE NAME (If any) 380/399							
DOSAGE FORM: Solution	STRENGTHS: 5%	ROUTE OF ADMINISTRATION: Topical							
(PROPOSED) INDICATION(S) FOR USE: To	regrow hair on the scalp.								
APPLICATION INFORMATION									
APPLICATION TYPE (check one) □ NEW DRUG APPLICATION (21 CFR 314.50)  □ BIOLOGICS LICENSE APPLICATION (21 CFR part 601)									
IF AN NDA, IDENTIFY THE APPROPRIATE TYPE   505 (b)(1)   505 (b)(2)									
IF AN ANDA, or 505(b)(2), IDENTIFY THE REI Name of Drug Rogaine (R)	FERENCE LISTED DRUG PRODUCT THAT IS THE BASIS FOR THE SUBMISSION Holder of Approved Application Pharmacia and Upjohn								
TYPE OF SUBMISSION (check one)	☐ ORIGINAL APPLICATION	☐ AMENDMENT TO A PENDING APPLICATION ☐ RESUBMISSION							
☐ PRESUBMISSION ☐ ANN	UAL REPORT ☐ ESTABLISHMENT DESCRIPTION SUPPLEMENT ☐ EFFICACY SUPPLEMENT								
☐ LABELING SUPPLEMENT	☐ CHEMISTRY MANUFACTURING AND								
IF A SUBMISSION OR PARTIAL APPLICATIO	N, PROVIDE LETTER DATE OF AGE	REEMENT TO PARTIAL SUBMISSION:							
IF A SUPPLEMENT, IDENTIFY THE APPROP	RIATE CATEGORY   CBE	☐ CBE-30 ☐ Prior Approval (PA)							
REASON FOR SUBMISSION Request for exemption from 21 CFR 201	I The second sec								
PROPOSED MARKETING STATUS (check on	e) PRESCRIPTION PRODUCT (Rx)	☑ OVER THE COUNTER PRODUCT (OTC)							
NUMBER OF VOLUMES SUBMITTED	1 THIS APPLICATION IS								
ESTABLISHMENT INFORMATION (Full establishment information should be provided in the body of the Application.)  Provide locations of all manufacturing, packaging and control sites for drug substance and drug product (continuation sheets may be used if necessary). Include name, address, contact, telephone number, registration number (CFN), DMF number, and manufacturing steps and/or type of testing (e.g., Final dosage form, Stability/testing) conducted at the site. Please indicate whether the site is ready for inspection or, if not, when it will be ready.									
		·							
Cross References (list related License Applications, INDs, NDAs, PMAs, 510(k)s, IDEs, BMFs, and DMFs referenced in the current application)									
NDA# 20-834									
,		- Annual Control of the Control of t							
FORM FDA 356h (4/00)	1 *	PAGE 1							

					ala de la filia de si				
This a	pplic	ation contains the following	items: (Chec	k all that apply)					
	1.	Index			1.4				
	2.	Labeling (check one)	Е	] Draft Labeling		☐ Final Prin	ted Labeling		
	3.	Summary (21 CFR 314.50(c))							
	4.	Chemistry section							
		A. Chemistry, manufact	turing, and controls information (e.g., 21 CFR 314.50(d)(1); 21 CFR 601.2)						
		B. Samples (21 CFR 31	4.50(e)(1); 21 CFR 601.2 (a)) (Submit only upon FDA's request)						
		C. Methods validation p	ackage (e.g., 21 CFR 314.50(e)(2)(i); 21 CFR 601.2)						
	5.	Nonclinical pharmacology and	toxicology section (e.g., 21 CFR 314.50(d)(2); 21 CFR 601.2)						
	6.	Human pharmacokinetics and	bioavailability section (e.g., 21 CFR 314.50(d)(3); 21 CFR 601.2)						
	7.	Clinical Microbiology (e.g., 21	CFR 314.50(d)(4))						
	8.	Clinical data section (e.g., 21	CFR 314.50(d)(5); 21 CFR 601.2)						
	9.	Safety update report (e.g., 21	CFR 314.50(d)(5)(vi)(b); 21 CFR 601.2)						
	10.	Statistical section (e.g., 21 CF	R 314.50(d)(6); 21 CFR 601.2)						
	11.	Case report tabulations (e.g.,	21 CFR 314.50(f)(1); 21 CFR 601.2)						
	12.	Case report forms (e.g., 21 CF	R 314.50(f)(2); 21 CFR 601.2)						
	13.	Patent information on any pate	ent which claims the drug (21 U.S.C. 355(b) or (c))						
	14.	A patent certification with resp	pect to any patent which claims the drug (21 U.S.C.355(b)(2) or (j)(2)(A)						
	15.	Establishment description (21	CFR Part 600, if applicable)						
	16.	Debarment certification (FD&C	Act 306(k)(1))						
	17.	Field copy certification (21 CF	R 314.50(k)(3))						
	18.	User Fee Cover Sheet (Form	FDA 3397)						
	19.	Financial Information (21 CFR	Part 54)						
$\boxtimes$	20.	OTHER (Specify) Application	for Exemption.						
CERTI	FICA	TION							
I agree to update this application with new safety information about the product that may reasonably affect the statement of contraindications, warnings, precautions, or adverse reactions in the draft labeling. I agree to submit safety update reports as provided for by regulation or as requested by FDA. If this application is approved, I agree to comply with all applicable laws and regulations that apply to approved applications, including, but not limited to the following:									
	2	2. Biological establishment star	ig practice regulations in 21 CFR Parts 210, 211or applicable regulations, Parts 606, and/or 820. Iment standards in 21 CFR Part 600. s in 21 CFR Parts 201, 606, 610, 660 and/or 809.						
	4. In the case of a prescription drug or biological product, prescription drug advertising regulations in 21 CFR 202.								
	<ol> <li>Regulations on making changes in application in FD&amp;C Act Section 506A, 21 CFR 314.71, 314.72, 314.97, 314.99, and 601.12.</li> <li>Regulations on Reports in 21 CFR 314.80, 314.81, 600.80 and 600.81.</li> </ol>								
If this a	7. Local, state and Federal environmental impact laws.  If this application applies to a drug product that FDA has proposed for scheduling under the Controlled Substances Act, I agree not to market the								
product until the Drug Enforcement Administration makes a final scheduling decision.  The data and information in this submission have been review and, to the best of my knowledge are certified to be true and accurate.									
The da   <b>Warni</b> r		d information in this submission A willfully false stateme	E .			certified to be true	e and accurate.		
SIGNAT	URE	OF RESPONSIBLE OFFICIAL OR A		TYPED NAME AND TITL Brian R. Schuster, Re	.E	nager	JUL 19 2001		
ADDRE	SS (S	treet, City, State, and ZIP Code)				TELEPHONE NUM			
515 Ea	asterr	Ave., Allegan,MI 49010				616-673-8451			
Public reporting burden for this collection of information is estimated to average 24 hours per response, including the time for reviewing instructions, searching existing data sources, gathering and maintaining the data needed, and completing and reviewing the collection of information. Send comments regarding this burden estimate or any other aspect of this collection of information, including suggestions for reducing this burden to:									
Food an CBER, 1401 R	nd Dru HFM- ockvil	le Pike	An agency may not conduct or sponsor, and a person is not required to respond to, a collection of information unless it displays a currently valid OMB control number.						
		) 20852-1448					DACE 2		